Background: HIV-infection is associated with premature decline of serum T. However, prevalence and biochemical characterization of hypogonadism in HIV-infected men are still to be well defined.

Method: Prospective, cross-sectional, observational study on HIV-infected men with ongoing Highly Active Antiretroviral Therapy (HAART). Serum TT, gonadotropins and sex hormone-binding globulin (SHBG) were measured by CI (Architect, Abbott, USA). TT was also assessed by a validated in house ID-LC-MS/MS. Free T (FT) was calculated by Vermeulen equation. Hypogonadism was defined as serum TT levels below ϰϮϬ ng/dL and/or free T levels below ϲϰ pg/ml.

Results: 315 consecutive HIV-infected men were enrolled (mean age 45.56±5.61 years; average duration of HIV-infection 16.57±10.45 years). Serum TT levels assessed by LC-MS/MS (mean 652.1±229.1 ng/dL) were significantly lower compared to CI (mean 740.2±274.7 ng/dL) (p<0.0001). As a consequence, prevalence of T deficiency was significantly higher comparing LC-MS-MS to CI (5.4% vs 3.2%, p<0.0001). 56 patients (17.8%) showed SHBG above the normal range (>71.4 nmol/L). Considering calculated FT, the prevalence of hypogonadism was 9.8% using LC-MS/MS and 7.0% using CI, with a significant difference between methodologies (p<0.0001). TT assessed with LC-MS-MS was directly related to TT assessed with CI (Beta=0.956, R²=0.913, p<0.0001), as well as FT (Beta=0.934, R²=0.873, p<0.0001). TT combined with luteinizing hormone (LH) levels was used to classify hypogonadism. By including compensated form of hypogonadism, the prevalence raised to ϰϱ.ϲй for TT and to ϰϳй for FT.

Conclusion: To the best of our knowledge, this is the first properly-designed prospective study aiming to investigate the gonadal status of HIV-infected men with both LC-MS/MS and CI, together with gonadotropins. Notwithstanding the strong correlation found between the two methodologies, the prevalence of hypogonadism results underestimated when CI is used compared to ID-LC-MS/MS in HIV-infected patients. In clinical practice, SHBG for calculated FT is essential for the detection of T deficiency, revealing the real prevalence of hypogonadism in this clinical setting.